Appl. No. 09/489,667 Reply to Office Action of October 24, 2005

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#### Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application:

#### Listing of Claims

#### 1-68. (Cancelled)

69. (Currently amended) An agent for treating pain comprising botulinum \_neurotoxin comprising a modified <del>clostridial</del> botulinum neurotoxin  $H_N$ , a botulinum neurotoxin L chain, and no functional Hc domain, wherein the modified clostridial neurotoxin is obtained by removing or modifying an HC domain of a native elostridial neurotoxin to form an intermediate clostridial neurotoxin and covalently coupling the modified botulinum neurotoxin being covalently coupled to substance P to the intermediate clostridial neurotoxin that the SO elostridial botulinum neurotoxin no longer binds to botulinum neurotoxin receptors at a neuromuscular junction with the same affinity as the native clostridial botulinum neurotoxin.

#### (Cancelled) 70.

- The agent of claim 69 wherein the (Currently amended) elestridial botulinum neurotoxin is a botulinum toxin selected from the group consisting of serotype A, serotype B, serotype  $C_1$ , serotype D, serotype E, serotype F and serotype G.
- (Currently amended) The agent of claim 69 wherein the 72. elestridial botulinum neurotoxin is botulinum toxin serotype A.

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- The agent of claim 69 wherein the (Currently amended) 73. elostridial botulinum-neurotoxin comprises an HN and an L chain agent is a fusion protein.
- (Cancelled) 74.
- The agent of elaim 73 wherein the L 75. (Currently amended) chain is obtained from an organism selected from the group consisting of Clostridial beratti, Clostridial butyricum, Clostridial betulinum, and Clostridial tetani claim 69, wherein the L chain is obtained from a botulinum toxin selected from the group consisting of botulinum toxin serotype A, serotype B, serotype Cl, serotype D, serotype E, serotype F and serotype G.
- The agent of claim 73 claim 69, (Currently amended) 76. wherein the  $H_{N}$  is obtained from a botulinum toxin selected from the group consisting of botulinum toxin serotype A, serotype B, serotype  $C_1$ , serotype D, serotype E, serotype F and serotype G.
- for treating agent (Previously presented) An comprising a botulinum toxin, without an  $H_{\rm c}$  that binds to receptors at the neuromuscular junction with the same affinity as native botulinum toxin, covalently coupled to substance P.
- for treating An agent (Previously presented) comprising a botulinum toxin serotype A, without a functional  $H_{\rm c}$ domain, covalently coupled to substance P.

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- for treating pain (Previously presented) An agent 79. comprising a botulinum toxin covalently coupled to substance P, wherein an  $H_c$  of the toxin has been removed.
- An agent for treating pain 80. (Previously presented) comprising a botulinum toxin serotype A covalently coupled to substance P, wherein an  $H_{\text{c}}$  of the toxin has been removed, the agent treats pain by acting on a projection neuron.